PCT

WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 5:

C12Q 1/68

(11) International Publication Number:

WO 90/04652

(43) International Publication Date:

3 May 1990 (03.05.90)

(21) International Application Number:

PCT/US89/04741

A1

(22) International Filing Date:

23 October 1989 (23.10.89)

(30) Priority data:

¥- _

₹

261,702

24 October 1988 (24.10.88) US

(71) Applicant: DNAX RESEARCH INSTITUTE OF MO-LECULAR AND CELLULAR BIOLOGY, INC. [US/ US]: 901 California Avenue, Palo Alto, CA 94304-1104

(US).

(72) Inventor: MACEVICZ, Stephen, C.; 1034 Innsbruck

Street, Livermore, CA 94550 (US).

(74) Common Representatives: MACEVICZ, Stephen, C. et al.; DNAX Research Institute of Molecular and Cellular Biology, Inc., 901 California Avenue, Palo Alto, CA 94304-1104 (US).

(81) Designated States: AT (European patent), BE (European patent), CH (European patent), DE (European patent), FR (European patent), GB (European patent), IT (European patent), JP, LU (European patent), NL (European patent), SE (European patent).

Published

With international search report.

Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(54) Title: DNA SEQUENCING BY MULTIPLE MIXED OLIGONUCLEOTIDE PROBES

(57) Abstract

A method is provided for sequencing nucleic acids without the need to separate similarly sized DNAs or RNAs by gel electrophoresis. The method relies on the separate hybridization of multiple mixed oligonucleotide probes to a target sequence. The mixed oligonucleotide probes comprise sequences of fixed and non-fixed bases corresponding to every possible permutation of fixed and non-fixed bases less than or equal to the length of the probes. For each probe, the hybridizations provide the number of times the probe's particular sequence of fixed bases appears in the target sequence. The target sequence is then mathematically reconstructed from this data and a knowledge of the probe sequences.

TARGET SEQUENCE PROBE 5'-CGAATGGAACTACCGTAACCT-3' 0 A 0 0 A 0 0 A 0 0 0 A 0 0 0 A 0 0 0 A 0 0 0 0 C 0 0 0 0000 CCOO 0 C C 0 0 0 C C G000 0G00 00G0 000G GG00 G 0 0 0 GGG 000 Ď 0 0 G0G0 G00G 0 G G 0 0 0 G G 0 0 G G T 0 0 0 0 0 T T